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NEWS 5 Feb 19 Access via Tymnet and SprintNet Eliminated Effective 3/31/02

NEWS 6 Mar 08 Gene Names now available in BIOSIS

NEWS 7 Mar 22 TOXLIT no longer available

NEWS 8 Mar 22 TRCTHERMO no longer available

NEWS 9 Mar 28 US Provisional Priorities searched with P in CA/CAplus and USPATFULL

NEWS 10 Mar 28 LIPINSKI/CALC added for property searching in REGISTRY

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NEWS 17 Apr 22 BIOSIS Gene Names now available in TOXCENTER

NEWS 18 Apr 22 Federal Research in Progress (FEDRIP) now available

NEWS 19 May 31 PCTFULL to be reloaded. File temporarily unavailable.

NEWS 20 Jun 03 New e-mail delivery for search results now available

NEWS EXPRESS February 1 CURRENT WINDOWS VERSION IS V6.0d, CURRENT MACINTOSH VERSION IS V6.0a(ENG) AND V6.0Ja(JP), AND CURRENT DISCOVER FILE IS DATED 05 FEBRUARY 2002

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=> s emulsion or multi-phas? or multiphas? or bi-phas? or biphas? or ((aqueous or water) and (lipid or oil or fat))

L1 655177 EMULSION OR MULTI-PHAS? OR MULTIPHAS? OR BI-PHAS? OR BIPHAS? OR ((AQUEOUS OR WATER) AND (LIPID OR OIL OR FAT))

=> s ribozyme or antisense or oligonucleotid? or aptamer

L2 231898 RIBOZYME OR ANTISENSE OR OLIGONUCLEOTID? OR APTAMER

=> s 11 and 12

L3 1117 L1 AND L2

=> s antioxidant or anti-oxid? and 13

L4 203311 ANTIOXIDANT OR ANTI-OXID? AND L3

=> s (antioxidant or anti-oxid?) and 13

L5 25 (ANTIOXIDANT OR ANTI-OXID?) AND L3

=> dup remove 15

PROCESSING COMPLETED FOR L5

19 DUP REMOVE L5 (6 DUPLICATES REMOVED)

=> s 16 and (anti-oxid? or antioxid) (5n) (ribozyme or antisense or oligonucleot? or aptamer)

L7 0 L6 AND (ANTI-OXID? OR ANTIOXID) (5N) (RIBOZYME OR ANTISENSE OR OLIGONUCLEOT? OR APTAMER)

=> d bib abs 16 1-19

L6 ANSWER 1 OF 19 CA COPYRIGHT 2002 ACS

AN 136:179054 CA

TI Transcription factor genes from Arabidopsis thaliana and their use for modifying plant traits

IN Pilgrim, Marsha; Creelman, Robert; Dubell, Arnold J.; Heard, Jacqueline;
Jiang, Cai-Zhong; Keddie, James; Adam, Luc; Ratcliff, Oliver; Reuber, J.
Lynne; Riechmann, Jose Luis; Yu, Guo-Liang; Pineda, Omaira

PA Mendel Biotechnology, Inc., USA

SO PCT Int. Appl., 941 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 8

PATENT NO. KIND DATE APPLICATION NO. DATE

```
PΙ
     WO 2002015675
                        A1
                             20020228
                                            WO 2001-US26189 20010822
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL,
              PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG,
              US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
              BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRAI US 2000-227439P
                       Ρ
                             20000822
     US 2000-713994
                       Α
                             20001116
     US 2001-837944
                       Α
                             20010416
AB
     The invention relates to 232 Arabidopsis plant transcription factor
     polypeptides, polynucleotides that encode them, homologs from a variety of
     plant species, and methods of using the polynucleotides and polypeptides
     to produce transgenic plants having advantageous properties compared to a
     ref. plant. Exemplary polynucleotides encoding the polypeptides of the
     invention were identified in the A. thaliana GenBank database using
     publicly available sequence anal. programs and parameters. Sequences
     initially identified were then further characterized to identify sequences
     comprising specified sequence strings corresponding to sequence motifs
     present in families of known transcription factors. Polynucleotide
     sequences meeting such criteria were confirmed as transcription factors.
     Further polynucleotides of the invention were identified by screening A.
     thaliana and/or other plant cDNA libraries with probes corresponding to
     known transcription factors under low stringency hybridization conditions.
     Addnl. sequences, including full-length coding sequences, were
     subsequently recovered by the rapid amplification of cDNA ends (RACE)
     procedure. The polynucleotides can be or were ectopically expressed in
     overexpressor or knockout plants and the changes in the characteristic(s)
     or trait(s) of the plant obsd.
              THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 2 OF 19 CA COPYRIGHT 2002 ACS
L6
ΑN
     136:149873 CA
TΙ
     IL-17 molecules and uses thereof
IN
     Medlock, Eugene; Yeh, Richard; Silbiger, Scott M.; Elliot, Gary S.;
     Nguyen, Hung Q.; Jing, Shuqian
PA
     Amgen, Inc., USA
so
     PCT Int. Appl., 242 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
FAN.CNT 3
     PATENT NO.
                      KIND DATE
                                            APPLICATION NO.
                                                             DATE
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                                            -----
ΡI
     WO 2002008285
                      A2
                            20020131
                                           WO 2001-US19861 20010621
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
             VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRAI US 2000-213125P
                       P
                            20000622
     US 2001-266159P
                       Ρ
                            20010202
     US 2001-810384
                       Α
                            20010316
AΒ
    Novel IL-17 like polypeptides and nucleic acid mols. encoding the same.
```

The invention also provides vectors, host cells, selective binding agents, and methods for producing IL-17 like polypeptides. Also provided for are methods for the treatment, diagnosis, amelioration, or prevention of diseases with IL-17 like polypeptides, agonists, or antagonists thereof.

- L6 ANSWER 3 OF 19 CA COPYRIGHT 2002 ACS
- AN 136:205405 CA
- TI Mixed micellar drug deliver system and method of preparation
- IN Modi, Pankai
- PA Generex Pharmaceuticals Incorporated, Can.
- SO U.S., 12 pp., Cont.-in-part of U.S. Ser. No. 386,285. CODEN: USXXAM
- DT Patent
- LA English
- FAN.CNT 5

tan.cm 3						
		PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	ΡI	US 6350458	B1	20020226	US 2000-543988	20000406
		US 6017545	Α	20000125	US 1998-21114	19980210
		US 6231882	B1	20010515	US 1998-216733	19981221
		US 6221378	В1	20010424	US 1999-386285	19990831
	PRAI	US 1998-21114	A2	19980210		
		US 1998-216733	A2	19981221		
		US 1999-386285	A2	19990831		

Pharmaceutical compns. comprising a macromol. pharmaceutical agent in AΒ micellar form are disclosed. The micelles are formed from an alkali metal alkyl sulfate, and at least one addnl. micelle-forming compd. as described in the specification. An alkali metal salicylate and a pharmaceutically acceptable edetate are also included in the compn. Micelle size ranges between about 1 and 10 nm. Methods for making and using the compns. are also disclosed. A buffer soln. was prepd. using 0.5 g sodium lauryl sulfate, 0.5 g sodium salicylate, and 0.25 g disodium edetate dissolved in 10 mL of water. The soln. was added to 16 mg (400 units) of insulin and mixed, to form micellar insulin. Sep., 100 mg of powd. Phosphatidylcholine-H was added to a glass beaker and to this powder was added 10 mL 50% ethanol. This soln. was then added to the above buffer soln., to give a 30 units/mg insulin soln., with vigorous mixing to form a mixed micellar soln. To this was added 0.6 mL of sodium hyaluronate and 0.2 mL of 2% menthol soln. contg. 3% sorbitol. Type II diabetic human volunteers took the micellar insulin orally. The oral insulin at a dosage of three times higher than the injected level, was comparable to the injected insulin.

RE.CNT 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L6 ANSWER 4 OF 19 CA COPYRIGHT 2002 ACS
- AN 136:252483 CA
- TI Clear oil-containing pharmaceutical compositions containing a therapeutic agent
- IN Chen, Feng-Jing; Patel, Mahesh V.; Fikstad, David T.
- PA USA
- SO U.S. Pat. Appl. Publ., 45 pp., Cont.-in-part of U.S. Ser. No. 751,968. CODEN: USXXCO
- DT Patent
- LA English
- FAN.CNT 4

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2002032171	A1	20020314	US 2001-877541	20010608
	us 6267985	B1	20010731	US 1999-345615	19990630
	US 6309663	B1	20011030	US 1999-375636	19990817
	US 2001024658	A1	20010927	US 2000-751968	20001229

```
PRAI US 1999-345615 A2 19990630
US 1999-375636 A2 19990817
US 2000-751968 A2 20001229
WO 2000-US18807 A 20000710
```

AB The present invention relates to pharmaceutical compns. and methods for improved solubilization of triglycerides and improved delivery of therapeutic agents. Compns. of the present invention include a carrier, where the carrier is formed from a combination of a triglyceride and at least 2 surfactants, at least one of which is hydrophilic. Upon diln. with an aq. medium, the carrier forms a clear, aq. dispersion of the triglyceride and surfactants. Thus, a formulation contained soybean oil, 80, Tween-20 200, and Tween-80 800 mg.

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L6 ANSWER 5 OF 19 CA COPYRIGHT 2002 ACS
```

AN 136:261833 CA

TI Sequence homologs of interleukin 17 and their use in diagnosis and treatment of immunol. diseases, inflammations and infections

IN Medlock, Eugene; Yeh, Richard; Silbiger, Scott M.; Elliott, Gary S.;
 Nguyen, Hung Q.; Jing, Shuqian

PA USA

SO U.S. Pat. Appl. Publ., 91 pp., Cont.-in-part of U.S. Ser. No. 810,384. CODEN: USXXCO

DT Patent

LA English

FAN.CNT 3

	PATENT NO.	KTND	DATE	APPLICATION NO.	DATE
ΡI	US 2002037524	A1	20020328	US 2001-886404	20010621
PRAI	US 2000-213125P	P	20000622		
	US 2001-266159P	P	20010202		
	US 2001-810384	A2	20010316		

AB Novel sequence homologs of IL-17 polypeptides (IL-17E) and nucleic acid mols. encoding the same are disclosed. The invention also provides vectors, host cells, antibodies and other selective binding agents, and methods for producing IL-17 like polypeptides. Also provided for are methods for the treatment, diagnosis, amelioration, or prevention of diseases with IL-17 like polypeptides, agonists, or antagonists thereof. Methods of high throughput drug screening for effectors of IL-17 polypeptides are another embodiment of the present invention.

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L6 ANSWER 6 OF 19 CA COPYRIGHT 2002 ACS
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AN 136:324075 CA

TI IL-17 receptor-like polypeptides, polynucleotides and antibodies for identification of agonists and antagonists and for diagnosis/treatment of immune diseases

IN Jing, Shuqian

PA USA

SO U.S. Pat. Appl. Publ., 54 pp., Cont.-in-part of U.S. Ser. No. 724,460. CODEN: USXXCO

DT Patent

LA English

FAN CNT 2

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI US 2002045213	A1	20020418	US 2001-809567	20010315
PRAI US 2000-189816P	P	20000316		
119 2000-724460	Δ2	20001128		

AB Disclosed are novel IL-17 receptor like polypeptides and nucleic acid mols. encoding the same. The invention also provides vectors, host cells, antibodies, antisense oligonucleotides, agonists and antagonists (including selective binding agents), and methods for producing IL-17 receptor like polypeptides. Also provided are methods for

the treatment, diagnosis, amelioration, or prevention of diseases assocd. with IL-17 receptor like polypeptides, e.g. immunol. diseases, autoimmune diseases, inflammation, transplant rejection, allergies, infections, obesity, anorexia, cachexia, neuronal diseases, lung diseases, skin diseases, kidney diseases, bone diseases, vascular diseases, cancer, etc. The invention further provides method for identifying antibody, small mol., protein, peptide, lipid, carbohydrate that mimicking or antagonizing the biol. activity of IL-17 receptor-like mol.

```
ANSWER 7 OF 19 CA COPYRIGHT 2002 ACS
L6
    136:11112 CA
ΑN
    Micellar pharmaceutical compositions for buccal and pulmonary application
ΤI
IN
    Modi, Pankaj
    Generex Pharmaceuticals Inc., Can.
PA
    PCT Int. Appl., 32 pp.
SO
    CODEN: PIXXD2
DT
     Patent
LA
    English
FAN.CNT 1
                                          APPLICATION NO. DATE
     PATENT NO.
                     KIND DATE
                                         _____
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                           -----
                                      WO 2001-CA661 20010507
    WO 2001087268 A1
                           20011122
PΙ
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            CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
            HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
            LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
            SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU,
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            BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRAI US 2000-574504
                    Α
                          20000519
     Pharmaceutical compns. comprising a macromol. pharmaceutical agent in
     mixed micellar form are disclosed. The mixed micelles are formed from an
     alkali metal alkyl sulfate, and at least three different micelle-forming
     compds. as described in the specification. Micelle size ranges between
     about 1 and 10 nm. A compn. contained powd. insulin, Na lauryl sulfate,
     deoxycholate, Na glycocholate, dibasic Na phosphate, and glycerin. A
     preferred method for administering the present compn. is through the
     buccal region of the mouth.
             THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 11
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 8 OF 19 CA COPYRIGHT 2002 ACS
L6
     135:376741 CA
AN
     Stable metal ion-lipid powdered pharmaceutical compositions
ΤI
     Dellamary, Luis A.; Riess, Jean; Schutt, Ernest G.; Weers, Jeffry G.;
IN
     Tarara, Thomas E.
     Alliance Pharmaceutical Corp., USA
PA
     PCT Int. Appl., 42 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LΑ
FAN.CNT 3
                                         APPLICATION NO. DATE
                     KIND DATE
     PATENT NO.
     _____
                                          WO 2001-US14824 20010508
                           20011115
                      A2
PΙ
     WO 2001085137
                     A3
                           20020418
     WO 2001085137
        W: AE, AG, AL, AM, AT, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH,
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             GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR,
             KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ,
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
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                           20000510
PRAI US 2000-568818
                      Α
    Microparticle compns. comprising metal ion-lipid complexes for
    drug delivery are described including methods of making the microparticle
    compns. and methods of treating certain conditions and disease states by
    administering the microparticle compns. The metal ion-lipid
    complexes can be combined with various drugs or active agents for
    therapeutic administration. The microparticle compns. of the present
     invention have superior stability to other microparticle compns. resulting
     in a microparticle compn. with longer shelf life and improved
    dispersibility. The microparticle compns. of the present invention have a
     transition temp. (Tm) of at least 20.degree. above the recommended storage
     temp. (Tst) for drug delivery. An aq. prepn. was prepd. by
    mixing two prepns., A and B, immediately prior to spray drying.
    prepn. A was comprised of a fluorocarbon-in-water
    emulsion in which 26 g perfluorooctyl bromide was dispersed in 33
     g water with the aid of 1.30 g of SPC-3 emulsifier (hydrogenated
     soy phosphatidylcholine). The prepn. B contained 0.162 g CaCl2.2H20 and
     0.162 g budesonide dissolved/suspended in 4 g water. The
     resulting microparticle of the sample had a PL-budesonide-CaCl2.2H20 wt.
     ratio of about 80:10:10. The mean vol. aerodynamic particle size of the
     dry powder was approx. 4.1 .mu.m.
     ANSWER 9 OF 19 CA COPYRIGHT 2002 ACS
L6
     135:271903 CA
AN
     IL-17 receptor like molecules and uses thereof
TI
     Jing, Shuqian; Medlock, Eugene; Yeh, Richard; Silbiger, Scott M.; Elliot,
IN
     Gary S.; Nguyen, Hung Q.
     Amgen Inc., USA
PA
     PCT Int. Appl., 239 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 3
                                           APPLICATION NO. DATE
                            \mathtt{DATE}
                      KIND
     PATENT NO.
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                                          WO 2001-US8688
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                            20010920
     WO 2001068705
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     US 2000-204208P
                      Ρ
                            20000512
     US 2000-723232
                       Α
                            20001127
                            20010202
     US 2001-266159P
                      Ρ
     Novel IL-17 receptor like polypeptides and nucleic acid mols. encoding the
            The invention also provides vectors, host cells, agonists and
     antagonists (including selective binding agents), and methods for
     producing IL-17 receptor like polypeptides. Also provided for are methods
     for treatment, diagnosis, amelioration, or prevention of diseases assocd.
     with IL-17 receptor like polypeptides, e.g. immune system dysfunction,
     inflammation, cancer and infection.
```

NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD,

RU, TJ, TM

```
AN
     134:198075
    Triglyceride-free compositions and methods for enhanced absorption of
TΙ
     hydrophilic therapeutic agents
     Patel, Mahesh V.; Chen, Feng-Jing
IN
     Lipocine, Inc., USA
PA
     PCT Int. Appl., 113 pp.
SO
     CODEN: PIXXD2
DΤ
     Patent
     English
LΑ
FAN.CNT 4
                                          APPLICATION NO. DATE
                     KIND DATE
     PATENT NO.
                           _____
                                         WO 2000-US18807 20000710
     WO 2001012155
                     A1
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                                        us 1999-375636
                      B1 20011030
     US 6309663
                           20020605
                                         EP 2000-947184
                                                           20000710
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     EP 1210063
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL
                                          US 2000-751968
                                                           20001229
                           20010927
     US 2001024658
                      A1
                           19990817
PRAI US 1999-375636
                      Α
                           20000710
     WO 2000-US18807
                      W
     The present invention relates to triglyceride-free pharmaceutical compns.,
     pharmaceutical systems, and methods for enhanced absorption of hydrophilic
     therapeutic agents. The compns. and systems include an absorption
     enhancing carrier, where the carrier is formed from a combination of at
     least two surfactants, at least one of which is hydrophilic. A
     hydrophilic therapeutic agent can be incorporated into the compn., or can
     be co-administered with the compn. as part of a pharmaceutical system.
     The invention also provides methods of treatment with hydrophilic
     therapeutic agents using these compns. and systems. For example, when a
     compn. contg. Cremophor RH40 0.30, Arlacel 186 0.20, Na taurocholate 0.18,
     and propylene glycol 0.32 g, resp., was used, the relative absorption of
     PEG 4000 as a model macromol. drug was enhanced by 991%.
             THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 1
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 11 OF 19 CA COPYRIGHT 2002 ACS
L6
     134:105846 CA
AN
     Clear aqueous dispersions of triglycerides and surfactants for
ΤI
     delivery of drugs and nutrients
     Chen, Feng-Jing; Patel, Mahesh V.
IN
     Lipocine, Inc., USA
PA
SO
     PCT Int. Appl., 103 pp.
     CODEN: PIXXD2
DΤ
     Patent
LA
     English
FAN.CNT 4
                                          APPLICATION NO. DATE
     PATENT NO.
                     KIND DATE
                                          _____
                                         WO 2000-US15133 20000602
                           20010111
                    A1
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     WO 2001001960
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             CU. CZ. DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
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ANSWER 10 OF 19 CA COPYRIGHT 2002 ACS

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ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD,
             SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA,
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     US 6267985
                            20010731
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     EP 1194120
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                            20020410
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         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
PRAI US 1999-345615
                            19990630
                       Α
     WO 2000-US15133
                       W
                            20000602
     The present invention relates to drug and nutrient delivery systems, and
AB
     in particular to pharmaceutical compns. and methods for improved
     solubilization of triglycerides and improved delivery of therapeutic
     agents. Compns. of the present invention include a triglyceride and a
     carrier, where the carrier is formed from a combination of at least two
     surfactants, at least one of which is hydrophilic. Upon diln. with an
     ag. solvent, the compn. forms a clear, ag. dispersion of
     the triglyceride and surfactants. An optional therapeutic agent can be
     incorporated into the compn., or can be co-administered with the compn.
     The invention also provides methods of enhancing triglyceride soly. and
     methods of treatment with therapeutic agents using these compns. Several
     formulations were presented of compns. that can be prepd. according to the
     present invention using a variety of therapeutic agents. Examples of
     ag. dispersions include: (1) Cremophor RH-40 0.75, Peceol 0.25,
     corn oil 0.40, and fenofibrate 0.10; (2) Cremophor RH-40 0.57,
     Crovol M-40 0.43, corn oil 0.40, and Rofecoxib 0.15; (3) Tween
     80 0.70, Tween 85 0.35, Miglyol 812 0.30, Paclitaxel 0.10, and PEG 400
     0.25; or (4) Kessco PEG 400 MO 0.33, corn oil 0.30, and
     Terbinafine 0.25 parts, resp.
RE.CNT 2
              THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L6
     ANSWER 12 OF 19 CA COPYRIGHT 2002 ACS
AN
     133:329593 CA
ΤT
     Low adenosine anti-sense oligonucleotide, compositions, kit and
     method for treatment of airway disorders associated with
     bronchoconstriction, lung inflammation, allergy(ies) and surfactant
     depletion
IN
     Nyce, Jonathan W.
     East Carolina University, USA
     PCT Int. Appl., 1592 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 1
                      KIND
                            DATE
     PATENT NO.
                                           APPLICATION NO.
                                                             DATE
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PΙ
    WO 2000062736
                      A2
                            20001026
                                           WO 2000-US8020
                                                             20000324
    WO 2000062736
                      A3
                            20011011
            AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP,
             KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,
            MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
            TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
             DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
             CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                            20010313
                                           BR 2000-6019
     BR 2000006019
                       Α
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EP 1168919 A2 20020109 EP 2000-919668 20000324
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO
PRAI US 1999-127958P P 19990406
WO 2000-US200008020W 20000324
WO 2000-US8020 W 20000324

OS MARPAT 133:329593

An in vivo method of selectively delivering a nucleic acid to a target AB gene or mRNA, comprises the topical administration, e.g. to the respiratory system, of a subject of a therapeutic amt. of an oligonucleotide (oligo) that is antisense to the initiation codon region, the coding region, the 5' or 3' intron-exon junctions or regions within 2 to 10 nucleotides of the junctions of the gene or antisense to a mRNA complementary to the gene in an amt. effective to reach the target polynucleotide and reducing or inhibiting expression. In addn. a method of treating an adenosine-mediated effect comprises topically administering to a subject an antisense oligo in an amt. effective to treat the respiratory, pulmonary, or airway disease. In order to minimize triggering adenosine receptors by their metab., the administered oligos have a low content of or are essentially free of adenosine. A pharmaceutical compn. and formulations comprise the oligo antisense to an adenosine receptor, genes and mRNAs encoding them, genomic and mRNA flanking regions, intron and exon borders and all regulatory and functionally related segments of the genes and mRNAs encoding the polypeptides, their salts and mixts. Various formulations contain a requisite carrier, and optionally other additives and biol. active agents. The low-adenosine or adenosine-free (des-A) agent for practicing the method of the invention may be prepd. by selecting a target gene(s), genomic flanking region(s), RNA(s) and/or polypeptide(s) assocd. with a disease(s) or condition(s) afflicting lung airways, obtaining the sequence of the mRNA(s) corresponding to the target gene(s) and/or genomic flanking region(s), and/or RNAs encoding the target $polypeptide(s)\,,\,\,selecting\,\,at\,\,least\,\,one\,\,segment\,\,of\,\,the\,\,mRNA\,\,which\,\,may\,\,be\,\,up$ to 60 % free of thymidine (T) and synthesizing one or more anti-sense oligonucleotide(s) to the mRNA segments which are free of adenosine (A) by substituting a universal base for A when present in the oligonucleotide. The agent may be prepd. by selection of target nucleic acid sequences with GC running stretches, which have low T content, and by optionally replacing A in the antisense oligonucleotides with a "Universal or alternative base". agent, compn. and formulations are used for prophylactic, preventive and therapeutic treatment of ailments assocd. with impaired respiration, lung allergy(ies) and/or inflammation and depletion lung surfactant or surfactant hypoprodn., such as pulmonary vasoconstriction, inflammation, allergies, allergic rhinitis, asthma, impeded respiration, lung pain, cystic fibrosis, bronchoconstriction. The present treatment is suitable for administration in combination with other treatments, e.g. before, during and after other treatments, including radiation, chemotherapy, antibody therapy and surgery, among others. Alternatively, the present agent is effectively administered prophylactically or therapeutically by itself for conditions without known therapies or as a substitute for therapies exhibiting undesirable side effects. The treatment of this invention may be administered directly into the respiratory system of a subject so that the agent has direct access to the lungs, or by other effective routes of administration, e.g. topically, transdermally, by implantation, etc., in an amt. effective to reduce or inhibit the symptoms of the ailment.

L6 ANSWER 13 OF 19 CA COPYRIGHT 2002 ACS

AN 132:203144 CA

TI Low-adenosine antisense oligonucleotide agents, compositions, kits and treatments for respiratory disorders

Nyce, Jonathan W. IN East Carolina University, USA PΑ SO PCT Int. Appl., 1343 pp. CODEN: PIXXD2 DΤ Patent English LΑ FAN.CNT 1 KIND DATE APPLICATION NO. DATE PATENT NO. ______ A2 WO 1999-US17712 19990803 WO 2000009525 20000224 PT 20000518 WO 2000009525 А3 W: AU, CA, CN, MX, RU, US RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE 20000306 AU 1999-53374 19990803 AU 9953374 Αl EP 1999-939006 EP 1102786 A2 20010530 19990803 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI 19980803 PRAI US 1998-95212P Ρ 19990803 WO 1999-US17712 MARPAT 132:203144 OS A compn. comprises a nucleic acid comprising an oligo antisense AB to a target such as polypeptide(s) assocd. With an ailment afflicting lung airways, genes and mRNAs encoding them, genomic and mRNA flanking regions, intron and exon borders and all regulatory and functionally related segments of the genes and mRNAs encoding the polypeptides, their salts and mixts. Various formulations contain a requisite carrier, and optionally other additives and biol. active agents. The agent of the invention may be prepd. by selecting a target gene(s), genomic flanking region(s), RNA(s) and/or polypeptide(s) assocd. with a disease(s) or condition(s) afflicting lung airways, obtaining the sequence of the mRNA(s) corresponding to the target gene(s) and/or genomic flanking region(s), and/or RNAs encoding the target polypeptide(s), selecting at least one segment of the mRNA which may be up to 60% free of thymidine (T) and synthesizing one or more antisense oligonucleotide(s) to the mRNA segments which are free of adenosine (A) by substituting a universal base for A when present in the oligonucleotide. The agent may be prepd. by selection of target nucleic acid sequences with GC running stretches, which have low T content, and by optionally replacing A in the antisense oligonucleotides with a universal base. The agent, compn. and formulations are used for prophylactic, preventive and therapeutic treatment of ailments assocd. with impaired respiration, allergy(ies) and/or inflammation, such as pulmonary vasoconstriction, inflammation, allergies, asthma, impeded respiration, lung pain, cystic fibrosis, bronchoconstriction, pulmonary hypertension and bronchoconstriction, chronic bronchitis, emphysema, chronic obstructive pulmonary disease (COPD), acute respiratory distress syndrome (ARDS), ischemic conditions including ischemia itself, and cancers such as leukemias, lymphomas, carcinomas, and the like, e.g. colon cancer, breast cancer, pancreatic cancer, lung cancer, hepatocellular carcinoma, kidney cancer, melanoma, hepatic metastasis, etc., as well as all types of cancers with may metastasize or have metastasized to the lung(s), including breast and prostate cancer. The present treatment is suitable for administration in combination with other treatments, e.g. before, during and after other treatments, including radiation, chemotherapy, antibody therapy and surgery, among others. The present agent is effectively administered preventatively, prophylactically or therapeutically by itself for conditions without known therapies, or as a substitute for, or in conjunction with, other therapies exhibiting undesirable side effects. The treatment of this invention may be

administered directly into the respiratory system of a subject, so that the agent has direct access to the airways and the lungs. The invention

is exemplified with specificity and pharmacokinetic studies using phosphorothioated antisense oligonucleotides targeted to the adenosine receptors Al, A2a, A2b, and A3.

- L6 ANSWER 14 OF 19 SCISEARCH COPYRIGHT 2002 ISI (R)
- AN 2000:476323 SCISEARCH
- GA The Genuine Article (R) Number: 326JE
- TI Adenosine A(1) receptor activation induces delayed preconditioning in rats mediated by manganese superoxide dismutase
- AU Dana A; Jonassen A K; Yamashita N; Yellon D M (Reprint)
- CS UNIV COLL LONDON HOSP, HATTER INST & CTR CARDIOL, LONDON WC1E 6DB, ENGLAND (Reprint); UNIV COLL LONDON HOSP, HATTER INST & CTR CARDIOL, LONDON WC1E 6DB, ENGLAND; UNIV COLL LONDON, SCH MED, LONDON WIN 8AA, ENGLAND
- CYA ENGLAND
- SO CIRCULATION, (20 JUN 2000) Vol. 101, No. 24, pp. 2841-2848.

 Publisher: LIPPINCOTT WILLIAMS & WILKINS, 530 WALNUT ST, PHILADELPHIA, PA 19106-3621.
 - ISSN: 0009-7322.
- DT Article; Journal
- FS LIFE; CLIN
- LA English
- REC Reference Count: 52
 - *ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS*
- AB Background-We have previously described a second window of protection against infarction in rabbits 24 to 72 hours after adenosine A, receptor (A(1)R) activation. In this study, we examined the potential role of the mitochondrial antioxidant manganese superoxide dismutase (Mn-SOD) as a potential end effector in mediating this protection.

Methods and Results-Rats were treated with an intravenous bolus of the AIR agonist 2-chloro-N-6-cyclopentyladenosine (CCPA, 75 mu g/kg) or saline vehicle. They were also given a 5 mg/kg IV infusion of a 22-mer phosphorothicate oligodeoxynucleotide (ODN) with sequence antisense to the initiation site of rat Mn-SOD mRNA. Sense ODN and scrambled ODN were used as controls. Twenty-four hours later, hearts were isolated and perfused with buffer at constant pressure and subjected to 35 minutes of regional ischemia and 2 hours of reperfusion. Treatment with CCPA compared with saline vehicle (control) significantly reduced infarct size, expressed as percentage of myocardium at risk (22.3+/-3.3% versus 42.1+/-3.8%, respectively; P=0.001). This protection was completely abolished by prior treatment with antisense ODN, which had no effect on its own. Neither sense ODN nor scrambled ODN had an effect on the CCPA-induced delayed cardioprotection. In separate animals, 24 hours after the same treatment, hearts were assayed for Mn-SOD content and activity. CCPA treatment induced a significant increase in myocardial Mn-SOD content and activity compared with the control condition; this increase was abolished by pretreatment with antisense ODN.

Conclusions-This is the first study to show that transient A(1)R activation induces delayed cardioprotection in the rat. These results strongly suggest an important role for mitochondrial Mn-SOD as a potential end effector of this protection.

- L6 ANSWER 15 OF 19 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE 1
- AN 2000:399577 BIOSIS
- DN PREV200000399577
- TI Reactive oxygen species stimulate p44/42 mitogen-activated protein kinase and induce p27Kipl: Role in angiotensin II-mediated hypertrophy of proximal tubular cells.
- AU Hannken, Tete; Schroeder, Regine; Zahner, Gunther; Stahl, Rolf A. K.; Wolf, Gunter (1)
- CS (1) Department of Medicine, Division of Nephrology and Osteology, University of Hamburg, University Hospital Eppendorf, Martinistrasse 52,

Pavilion 61, D-20246, Hamburg Germany

SO Journal of the American Society of Nephrology, (August, 2000) Vol. 11, No. 8, pp. 1387-1397. print. ISSN: 1046-6673.

- DT Article
- LA English
- SL English
- Angiotensin II (AngII) induces G1 phase arrest and hypertrophy of cultured AΒ renal proximal tubular cells. In previous studies, it was shown that these effects depend on oxygen radical-mediated induction of p27Kip1, an inhibitor of cyclin-dependent kinases. The present study was undertaken to investigate whether mitogen-activated protein (MAP) kinases serve as signaling intermediates between AngII-induced oxidative stress and induction of p27Kipl. AngII (10-7 M) induces a biphasic phosphorylation pattern of p44/42 MAP kinase with an early phosphorylation after 2 min and a later, second phosphorylation peak after prolong incubation (12 h) in cultured proximal tubular cells from two different species (MCT and LLC-PK1 cells). Total protein expression of MAP kinase was not changed by AngII. These phosphorylation patterns of p44/42 MAP kinase caused activation of the enzyme, as detected by phosphorylated MAP substrate Elk-1 after immunoprecipitation of MAP kinase. Exogenous H2O2 also stimulates a biphasic phosphorylation of p44/42 MAP kinase. The flavoprotein inhibitor diphenylene iodinium, as well as the anti-oxidant N-acetylcysteine, prevented AngII-induced p44/42 MAP kinase phosphorylation, indicating involvement of reactive oxygen species generated by membrane-bound NAD(P)H oxidase. The MAP kinase kinase inhibitor PD98059 completely inhibits AngII-induced p27Kip1 expression and 3(H)leucine incorporation into proteins as a previously established marker of cell hypertrophy. PD98059 did not attenuate AngII-stimulated intracellular synthesis of oxygen radicals. Transient transfection with p44/42 MAP kinase antisense, but not sense, phosphorothicate-modified oligonucleotides also prevented AngII-induced MAP kinase phosphorylation, p27Kip1 expression, and cell hypertrophy. Furthermore, induction of p27Kip1 by H2O2 was also abolished in the presence of PD98059. Although AngII induces phosphorylation of the stress-activated p38 MAP kinase, inhibition of this enzyme with SB203580 failed to attenuate induced P27Kip1 expression and hypertrophy. These data provide evidence that AngII-mediated oxygen stress leads to the phosphorylation of p44/42 MAP kinase in proximal tubular cells. Activation of this enzyme is essential for p27Kip1 expression, G1 phase arrest, and hypertrophy of proximal tubular cells. These findings may lead to new concepts concerning interference of the development of proximal tubular hypertrophy, which may eventually turn into a maladaptive process in vivo leading ultimately to tubular atrophy and tubulointerstitial fibrosis.
- L6 ANSWER 16 OF 19 CA COPYRIGHT 2002 ACS
- AN 132:54836 CA
- TI Adenosine receptor-modulator compositions and methods for prevention and treatment of cardiopulmonary and renal failure or damage associated with ischemia, endotoxin release, ARDS, or brought about by administration of certain drugs
- IN Nyce, Jonathan W.; Hill, Jeffrey L.
- PA Epigenesis Pharmaceuticals, Inc., USA
- SO PCT Int. Appl., 252 pp.
- CODEN: PIXXD2
- DT Patent
- LA English
- FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9963938	A2	19991216	WO 1999-US12775	19990608
	WO 9963938	A3	20000127		

W: AU, CA, CN, MX, US RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE CA 1999-2316994 19990608 19991216 CA 2316994 AΑ 20000628 EP 1999-930160 19990608 EP 1011608 A2 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, NL, SE, PT, IE, FI BR 2000-1593 20000412 20011204 BR 2000001593 Α Р 19980608 PRAI US 1998-88501P US 1998-88657P P 19980609 19980609 US 1998-93972 Α WO 1999-US12775 19990608 MARPAT 132:54836 OS

A pharmaceutical compn. comprises an agent such as an adenosine A2a AB agonist agent and/or nucleic acid comprising an oligonucleotide (oligo) that is anti-sense to an adenosine A1, A2a, A2b or A3 receptor gene, mRNA, flanking regions or regions bridging the intron/exon borders, which oligos are effective to prevent, alleviate or inhibit adenosine-mediated cardiac, pulmonary and/or renal functional difficulties, damage or failure, such as those obsd. in diseases and conditions such as ARDS, hypoxia, etc. or assocd. with the administration of therapeutic and diagnostic agents such as adenosine cisplatin, metal ion-contg. agents, etc., mixts. thereof, and optionally a surfactant, a carrier and other therapeutic and diagnostic agents and other formulation components. The compn. is provided in the form of various formulations that are, for example, effective for preventing or alleviating bronchoconstriction, allergy and/or inflammation assocd. with ARDS, RDS, etc., deleterious side effects obsd. upon treatment of SVT patients, upon administration of cardiac stress tests or imaging tests, etc.

- L6 ANSWER 17 OF 19 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE
- AN 1999:330249 BIOSIS
- DN PREV199900330249
- TI Exercise provides direct **biphasic** cardioprotection via manganese superoxide dismutase activation.
- AU Yamashita, Nobushige; Hoshida, Shiro (1); Otsu, Kinya; Asahi, Michio; Kuzuya, Tsunehiko; Hori, Masatsugu
- CS (1) Cardiovascular Division, Osaka Rosai Hospital, 1179-3 Nagasonecho, Sakai, Osaka, 591-8025 Japan
- SO Journal of Experimental Medicine, (June 11, 1999) Vol. 189, No. 11, pp. 1699-1706.
 ISSN: 0022-1007.
- DT Article
- LA English
- SL English
- Epidemiologic investigations have shown that exercise reduces morbidity AB and mortality from coronary artery disease. In this study, using a rat model, we attempted to determine whether exercise can reduce ischemic injury to the heart and elucidate a mechanism for the cardioprotective effect of exercise. Results showed that exercise significantly reduced the magnitude of a myocardial infarction in biphasic manner. The time course for cardioprotection resembled that of the change in manganese superoxide dismutase (Mn-SOD) activity. The administration of the antisense oligodeoxyribonucleotide to Mn-SOD abolished the expected decrease in infarct size. We showed that the level of tumor necrosis factor alpha (TNF-alpha) and interleukin lbeta (IL-lbeta) increased after exercise. The simultaneous administration of the neutralizing antibodies to the cytokines abolished the exercise-induced cardioprotection and the activation of Mn-SOD. Furthermore, TNF-alpha can mimic the biphasic pattern of cardioprotectionand activation of Mn-SOD. An antioxidant completely abolished cardioprotection and the activation of Mn-SOD by exercise or the injection of TNF-alpha as well

as exercise-induced increase in TNF-alpha and IL-lbeta. The production of reactive oxygen species and endogenous TNF-alpha and IL-lbeta induced by exercise leads to the activation of Mn-SOD, which plays major roles in the acquisition of biphasic cardioprotection against ischemia/reperfusion injury in rats.

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L6
    ANSWER 18 OF 19 CA COPYRIGHT 2002 ACS
AN
    128:275101 CA
ΤI
    Gas and gaseous precursor filled microspheres as topical and subcutaneous
    delivery vehicles
IN
    Unger, Evan C.; Matsunaga, Terry O.; Yellowhair, David
PA
    Imarx Pharmaceutical Corp., USA
SO
    U.S., 40 pp., Cont.-in-part of U.S. Ser. No. 307,305.
    CODEN: USXXAM
DT
    Patent
    English
LΑ
FAN.CNT 19
     PATENT NO.
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PRAI US 1989-455707

В2

19891222

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US 1990-569828
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US 1991-716899
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US 1991-717084
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US 1993-76250
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US 1993-159687
                A2 19931130
US 1993-160232
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US 1994-307305
                A2 19940916
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WO 1990-US7500
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US 1991-750877 A3 19910826
US 1992-818069 A3 19920108
WO 1992-US2615 A 19920331
US 1992-967974
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US 1993-17683
                 A3
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US 1993-18112
                 В3
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US 1993-85608
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WO 1994-US13817 W
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US 1995-395683
                A3
                       19950228
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Gas and gaseous precursor filled microspheres, and foams provide novel
topical and s.c. delivery vehicles for various active ingredients,
including drugs and cosmetics. Gas and gaseous precursor filled
microcapsules were prepd. from dipalmitoylphosphatidylcholine.
ANSWER 19 OF 19
                    MEDLINE
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20091286 PubMed ID: 10099076
Carbonyl-trapping therapeutic strategies.
Shapiro H K
Department of Pharmacology, Temple University Medical School,
Philadelphia, PA, USA.
AMERICAN JOURNAL OF THERAPEUTICS, (1998 Sep) 5 (5) 323-53. Ref: 273
Journal code: DB7; 9441347. ISSN: 1075-2765.
United States
Journal; Article; (JOURNAL ARTICLE)
General Review; (REVIEW)
(REVIEW, TUTORIAL)
English
Priority Journals
200001
Entered STN: 20000204
Last Updated on STN: 20000204
Entered Medline: 20000124
Under conditions of oxidative stress, aldehyde or ketone products are
generated nonenzymatically by lipid peroxidation or form
spontaneously from simple sugars. Many aldehydes, in particular, are
cytotoxic. They may react with primary amine groups to form Schiff bases,
which may subsequently rearrange into more chemically stable structures.
In biological systems, such reactions may disrupt normal
oligonucleotide structure, may interfere with the biological
activity of numerous structural or enzymatic polypeptides, and may
covalently cross-link proteins and lipids (eg,
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phosphatidylethanolamine). Once thought to be largely epiphenomenal, such

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events are now known to be central to the etiologies of a spectrum of neurodegenerative diseases, chronic inflammatory diseases, and pathophysiologically related disorders. Opportunities exist for therapeutic intervention in these disease states by use of certain water-soluble, small-molecular-weight drugs that contain primary amine groups. Such pharmaceutical agents, administered orally, can form Schiff-base derivatives with toxic carbonyl substances and thus protect cellular components. Future studies of such carbonyl-trapping agents may include their use in combination with other classes of drugs, such as antioxidants, anti-inflammatory products, or neuroactive agents. This conceptually simple approach may offer new opportunities for improved clinical management of many chronic disease states.

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